Date of Deposit hereby certify that this paper or fee is being deposited with the United States Postal Service "Express Mail Post Office Addressee" service under 37 CFR 1.10 on the date adicated abortoglative Edderson Residence (FR persons are required)	II S Parent and	PTO/SB/30 (01-03) Approved for use through 04/30/2003. OMB 0651/0031 Frademark Office; U.S. DEPARTMENT OF COMMERCE formation unless it contains a valid OMB control furniber.
P.O. Box 1450, Alexandria 422131450	Application Number	09/541,848
For (DOF)	Filing Date	April 3, 2000
Continued Examination (RCE)	First Named Inventor	Chen
Transmittal Address to: Commissioner for Patents P. O. Box 1450, Mail Stop RCE Alexandria, VA 22313-1450	Art Unit	1635
	Examiner Name	Schultz, James
	Attorney Docket Numb	per HYB-015US4 (1006/016)
This is a Request for Continued Examination (RCE) under 37 CFR 1.114 of the above-identified application. Request for Continued Examination (RCE) practice under 37 CFR 1.114 does not apply to any utility or plant application filed prior to June 8, 1995, or to any design application. See Instruction Sheet for RCEs (not to be submitted to the USPTO) on page 2. 1. Submission required under 37 CFR 1.114		
a. Previously submitted		
i. Consider the amendment(s)/reply under 37 CFR 1.116 previously filed on May 2, 2003 (copy enclosed) (Any unentered amendment(s) referred to above will be entered).		
li. Consider the arguments in the Appeal Brief or Rely Brief previously filed on		
lii.		
b Enclosed I Amendment/Reply ii Affidavit(s)/ Declaration(s)	iii.	ation Disclosure Statement (IDS)
2. Miscellaneous Suspension of action on the above-identified application is requested under 37 CFR 1.103(c) for a a. period of months. (Period of suspension shall not exceed 3 months; Fee under 37 CFR 1.17(i) required) b. Other		
3. Fees The RCE fee under 37 CFR 1.17(e) is required by 37 CFR 1.114 when the RCE is filed. The Director is hereby authorized to charge the following fees, or credit any overpayments, to Deposit Account No. 50-2285		
i. RCE fee required under 37 CFR 1.17(e)	08/20/	2003 MBIZUNES 00000063 09541848
ii. Extension of time fee (37 CFR 1.136 and 1	.17) 01 FC:	2801 375.00 QP
iii Other b. ✓ Check in the amount of \$ 580.00	enclos	ed
c. Payment by credit card (Form PTO-2038 enclosed)		
WARNING: Information on this form may become public. Credit card information should not be included on this form. Provide credit card information and authorization on PTO-2038.		
SIGNATURE OF APPLICANT, ATTORNEY, OR AGENT REQUIRED		
Name (Print/Type) Wayne A. Keown, Ph.D. Signature	Reg Date	istration No. (Attorney/Agent) 33,923
CERTIFICATE OF MAILING		
I hereby certify that this correspondence is being deposited with the United States Postal Service with sufficient postage as Express Mail in an envelope addressed to: Commissioner For Patents, Mail Stop RCE. P.O. Box 1450, Alexandria, VA 22313-1450 on the date shown below.		
Name (Print/Type) Me a sile Life are Signature Date S/18/83		

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Page I of 2

This collection of information is required by 37 CFR 1.114. The information is required to obtain or retain a benefit by the public which is to file (and by the USPTO to process) an application. Confidentiality is governed by 35 U.S.C. 122 and 37 CFR 1.14. This collection is estimated to take 12 minutes to complete, including gathering, preparing, and submitting the completed application form to the USPTO. Time will vary depending upon the individual case. Any comments on the amount of time you require to complete this form and/or suggestions for reducing this burden, should be sent to the Chief Information Officer, U.S. Patent and Trademark Office, U.S. Department of Commerce, Washington, DC 20231. DO NOT SEND FEES OR COMPLETED FORMS TO THIS ADDRESS. SEND TO:

Commissioner for Patents, Box RCE, Washington, DC 20231.

If you need assistance in completion the form cell 4 ROS CTO CASE COMPLETED.



I THE UNITED STATES PATENT AND TRADEMARK OFFICE

IN APPLICATION OF:

ART UNIT: 1635

CHEN et al.

EXAMINER: Wang, Andrew J.

SERIAL NO.: 09/541,848

FILED: April 3, 200

FOR: MDM2 SPECIFIC ANTISENES

OLIGONUCLEOTIDES

CERTIFICATE OF FIRST CLASS MAILING UNDER 37 C.F.R. § 1.8

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Date: 4/22/03

Signature Ulusa A. Sumpton

Printed Name: Melissa A. Simpson

Box AF

Assistant Commissioner for Patents Washington, DC 20231

AMENDMENT AND RESPONSE UNDER 37 C.F.R. § 1.116

Dear Sir:

In response to the Final Office Action dated October 22, 2002, please reconsider the above-referenced patent application in view of the following amendment and remarks. Included with this response is a Request for an Extension of Time for 3 months under 37 C.F.R. § 1.136(a) and the appropriate fee.

"Express Mail" mailing label number (28494570).

Date of Deposit 418709

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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

<u>AMENDMENTS</u>

LISTING OF THE CLAIMS

- 1. (Presently Amended) A method of inhibiting expression of MDM2 in a mammal, the method comprising administering to the mammal an effective MDM2-expression inhibiting amount of an anti-MDM2 antisense oligonucleotide, wherein said antisense oligonucleotide comprises from about 8 to about 50 nucleotides that inhibits MDM2 protein expression, said oligonucleotide binding to mdm2-encoding RNA and being complementary to a sequence that overlaps by at least one nucleotide a sequence within the mdm2 RNA, which sequence within the mdm2 RNA is selected from the group consisting of SEQ ID NOS: 2, 3, 4, 7, 8, 9, 10, 11, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, and 24.
- The method according to claim 1 comprising co-administering a cancer chemotherapeutic agent.
- 3. The method according to claim 2, wherein the cancer therapeutic agent is 10-hydorxycamptothecin, adriamycin, or 5-fluorouracil.
- 4. The method according to claim 1 comprising co-treating the mammal with anti-cancer levels of radiation.
- 5. (Presently Amended) A method of inhibiting cancer *in vivo*, the method comprising administering a cancer-inhibiting amount of an anti-MDM2 antisense oligonucleotide, wherein the cancer involves over expression of MDM2, wherein said antisense oligonucleotide comprises from about 8 to about 50 nucleotides that inhibits MDM2 protein expression, said oligonucleotide binding to mdm2-encoding RNA and being complementary to a sequence that overlaps by at least one nucleotide a sequence within the mdm2 RNA, which sequence within the mdm2 RNA is selected from the